

AMENDMENT TO THE CLAIMS

1-50. (Canceled)

51. (Withdrawn) A composition, for delivery of a therapeutic agent to a neuronal cell, comprising:

a therapeutic agent which inhibits at least one member of the Rho group of GTPases, and

a neuronal cell targeting component, which component comprises a Hc domain of botulinum C1 toxin, or a fragment thereof which retains the function of the native Hc domain,

wherein the Hc domain has been made recombinantly.

52. (Currently amended) A composition, for delivery of a therapeutic agent to a neuronal cell, comprising:

a therapeutic agent which inhibits at least one member of the Rho group of GTPases,

a neuronal cell targeting component, which component comprises a Hc domain of botulinum C1 toxin, or a fragment thereof which retains the function of the native Hc domain, and

a domain for translocation of the therapeutic agent into a cell,

wherein the Hc domain has been made recombinantly, and

the therapeutic agent is an ADP-ribosyltransferase

A composition according to claim 51 further comprising a domain for translocation of the therapeutic agent into a cell.

53. (Currently amended) A composition according to claim 52, wherein the translocation domain is derived from a clostridial source.

54. (Withdrawn – Currently amended) A composition according to claim 52, wherein the translocation domain is derived from a non-clostridial source.

55. (Currently amended) A composition according to claim 53, wherein the translocation domain is derived from *C. botulinum*, *C. butylicum*, *C. argentinense* or *C. tetani*.

56. (Withdrawn – Currently amended) A composition according to claim 54, wherein the translocation domain comprises a translocation domain of diphtheria toxin, Pseudomonas exotoxin A, influenza virus haemagglutinin fusogenic peptides or amphiphilic peptides.

57. (Original) A composition according to claim 52, wherein the translocation domain comprises a member selected from the group consisting of botulinum C1 toxin and fragments thereof, and diphtheria toxin and fragments thereof.

58. (Original) A composition according to claim 52 wherein the translocation domain is a membrane disrupting peptide.

59. (Withdrawn – Currently amended) A composition according to claim [[51]] 52, wherein the therapeutic agent is selected from the group consisting of drugs, growth factors, enzymes, DNA, modified viruses, drug release systems, and a combination thereof.

60. (Withdrawn – Currently amended) A composition according to claim [[51]] 52, wherein the therapeutic agent is a C3 enzyme.

61. (Withdrawn) A composition according to claim 60, wherein the C3 enzyme is derived from *C. botulinum*, *C. limosum*, *B. cereus*, *S. aureus*, *C. acetobutylicum*, *S. pyogenes*, *L. monocytogenes*.

62. (Withdrawn) A composition according to claim 60, wherein the C3 enzyme is selected from the group consisting of C3Stau2, C3Stau1, and C3bot.

63. (Withdrawn) A composition according to claim 60, wherein the C3 enzyme has an amino acid sequence selected from the group consisting of SEQ ID Nos: 1-10.

64. (Withdrawn – Currently amended) A composition according to claim [[51]] 52, wherein the therapeutic agent and the Hc domain are joined to each other directly or via a linker molecule.

65. (Original) A composition according to claim 52, wherein the therapeutic agent, the Hc domain and the translocation domain are joined to each other directly or via a linker molecule.

66. (Withdrawn – Currently amended) A composition according to claim 64, wherein the linker molecule is selected from the group consisting of [[[GGGGS)2, (GGGGS)3,]] the interdomain linker of cellulase, [[PPPIEGR,]] collagen[[-like]] spacer, trypsin-sensitive diphtheria toxin peptide, and linker molecules having an amino acid sequence of SEQ ID Nos: [[16-24]] 16-27.

67. (Currently amended) A composition according to claim 65, wherein the linker molecule is selected from the group consisting of [[[GGGGS)2, (GGGGS)3,]] the interdomain linker of cellulase, [[PPPIEGR,]] collagen[[-like]] spacer, trypsin-sensitive diphtheria toxin peptide, and linker molecules having an amino acid sequence of SEQ ID Nos: [[16-24]] 16-27.

68. (Withdrawn – Currently amended) A composition according to claim [[51]] 52, wherein the composition is a single polypeptide.

69. (Withdrawn – Currently amended) A composition according to claim [[51]] 52, wherein the composition is a dichain polypeptide.

70. (Withdrawn – Currently amended) A composition according to claim [[51]] 52, wherein the composition is a suspension, emulsion, solution or a freeze-dried powder.

71. (Withdrawn – Currently amended) A composition according to claim [[51]] 52, further comprising a pharmaceutically acceptable liquid.

72. (Withdrawn – Currently amended) A method of making a composition according to claim [[51]] 52, comprising expressing a DNA encoding the therapeutic agent and the neuronal cell targeting domain.